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E11
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L2
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YOU HAVE REQUESTED DATA FROM 3 ANSWERS - CONTINUE? Y/(N):y
L2
    ANSWER 1 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
    ΑN
    144:410795
DN
ΤI
    Recombinant Mycobacterium BCG adjuvant in vaccination
    Laeufer, Albrecht; Eisele, Bernd; ***Grode, Leander***
PA
    Vakzine Projekt Management G.m.b.H., Germany
SO
    Eur. Pat. Appl., 17 pp.
    CODEN: EPXXDW
DT
    Patent
LA
    English
FAN.CNT 1
                      KIND DATE APPLICATION NO. DATE
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                              20060426 EP 2004-25096
    EP 1649869
                        A1
                                                               20041021
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            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ,
            NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG,
            SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN,
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        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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            CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
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    CN 101048178
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IN 2007DN02871 A 20070817 IN 2007-DN2871 MX 200704734 A 20070713 MX 2007-4734 KR 2007068398 A 20070629 KR 2007-709076
                                                                   20070418
                                                                   20070419
                                                               20070420
PRAI EP 2004-25096
     EP 2004-25096 A
WO 2005-EP11127 W
                              20041021
20051016
AB
     The authors disclose the use of ***urease*** -deficient Mycobacterium
     BCG expressing listeriolysin as an adjuvant in vaccination. In one
     example, a tumor vaccine comprises a allogeneic prostate carcinoma cell
     line, transgenic for interferon-.gamma. and interleukin-2, in combination
     with the foregoing bacterial cell adjuvant.
RE.CNT 6
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
ΙN
     Laeufer, Albrecht; Eisele, Bernd; ***Grode, Leander***
AΒ
     The authors disclose the use of ***urease*** -deficient Mycobacterium
     BCG expressing listeriolysin as an adjuvant in vaccination. In one
     example, a tumor vaccine comprises a allogeneic prostate carcinoma. . .
ΙT
    Vaccines
        (antimalarial;
                       ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as adjuvant for)
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (autoantigens, microbial; ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as adjuvant in vaccination against)
ΤТ
     Prostate gland, neoplasm
        (carcinoma; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as vaccine adjuvant for cytokine-transgenic cell
        immunogens)
ΙT
     Intestine, neoplasm
        (colon, carcinoma;
                            ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
        cell immunogens)
    Carcinoma
ΤТ
        (colon; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as vaccine adjuvant for cytokine-transgenic cell
        immunogens)
ΤТ
     Carcinoma
        (head and neck squamous cell carcinoma; ***urease*** -deficient
        Mycobacterium BCG expressing listeriolysin as vaccine adjuvant for
        cytokine-transgenic cell immunogens)
     Cell adhesion molecules
ΙT
     Interleukin 12
     Interleukin 2
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (in combination with
                              ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as adjuvant in vaccination)
ΙT
     Hemolysins
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (listeriolysins O; ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as adjuvant in vaccination)
ΙT
    Antigens
     Tumor antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (microbial; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as adjuvant in vaccination against)
ΙT
     Lung, neoplasm
```

```
(non-small-cell carcinoma; ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
        cell immunogens)
ΙT
    Lysosome
                        ***urease*** -deficient Mycobacterium BCG expressing
        (phagolysosome;
        listeriolysin as adjuvant in vaccination in relation to)
ΤТ
        (prostatic; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as vaccine adjuvant for cytokine-transgenic cell
        immunogens)
ΙT
    Carcinoma
        (pulmonary non-small-cell; ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
        cell immunogens)
ΤТ
    Kidney, neoplasm
        (renal cell carcinoma;
                                ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
        cell immunogens)
    Carcinoma
ΤТ
                     ***urease*** -deficient Mycobacterium BCG expressing
        (renal cell;
        listeriolysin as vaccine adjuvant for cytokine-transgenic cell
        immunogens)
    Head and Neck, neoplasm
ΙT
        (squamous cell carcinoma; ***urease*** -deficient Mycobacterium BCG
        expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
       cell immunogens)
ΙT
    Vaccines
        (tumor; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as adjuvant for)
ΙT
    MSP-1 (protein)
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        ( ***urease*** -deficient Mycobacterium BCG expressing listeriolysin
        as adjuvant for)
ΙT
     Plasmodium falciparum
        ( ***urease*** -deficient Mycobacterium BCG expressing listeriolysin
        as adjuvant for merozoite surface protein of)
ΙT
    Malaria
        ( ***urease*** -deficient Mycobacterium BCG expressing listeriolysin
        as adjuvant for vaccination against)
ΙT
    Mycobacterium BCG
     Mycobacterium bovis
        ( ***urease*** -deficient Mycobacterium BCG expressing listeriolysin
        as adjuvant in vaccination)
ΤТ
    Antigen-presenting cell
     Brain, neoplasm
     Dendritic cell
     Mammary gland, neoplasm
     Melanoma
     Neoplasm
        ( ***urease*** -deficient Mycobacterium BCG expressing listeriolysin
        as vaccine adjuvant for cytokine-transgenic cell immunogens)
TΤ
    Antimalarials
     Antitumor agents
        (vaccines; ***urease*** -deficient Mycobacterium BCG expressing
        listeriolysin as adjuvant for)
```

IT Interferons

RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(.gamma.; in combination with ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 884349-82-0

RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (amino acid sequence; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 9002-13-5D, ***Urease*** , subunit C

RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)

(deficiency; ***urease*** -deficient Mycobacterium BCG expressing listeriolysin as adjuvant in vaccination)

IT 884349-81-9, DNA (Listeria monocytogenes gene hyl)
RL: BSU (Biological study, unclassified); PRP (Properties); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
(nucleotide sequence; ***urease*** -deficient Mycobacterium BCG
expressing listeriolysin as adjuvant in vaccination)

- L2 ANSWER 2 OF 3 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>
- DN PREV200510303569
- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU ***Grode, Leander*** ; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; Kaufmann, Stefan H. E. [Reprint Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany Kaufmann@mpiib-Berlin.mpg.de
- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.

 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 23 Nov 2005
 Last Updated on STN: 23 Nov 2005
- AB The tuberculosis vaccine Mycobacterium bovis bacille Calmette-Guerin (BCG) was equipped with the membrane-perforating listeriolysin (Hly) of Listeria monocytogenes, which was shown to improve protection against Mycobacterium tuberculosis. Following aerosol challenge, the Hly-secreting recombinant BCG (hly(+) rBCG) vaccine was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG strain. The isogenic, ***urease*** C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for Hly activity, exhibited still higher vaccine efficacy than parental BCG. Delta ureC hly(+) rBCG also induced profound protection against a member of the M. tuberculosis Beijing/W genotype family while parental BCG failed to do so consistently. Hly not only promoted antigen translocation into the cytoplasm but also apoptosis of infected macrophages. We concluded that superior vaccine efficacy of

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Delta ureC hly(+) rBCG as compared with parental BCG is primarily based on
     improved cross-priming, which causes enhanced T cell-mediated immunity.
ΑU
       ***Grode, Leander*** ; Seiler, Peter; Baumann, Sven; Hess, Juergen;
     Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian;
    Bandermann, Silke; Smith, Debbie; . . .
     . . was shown to protect significantly better against aerosol infection
    with M. tuberculosis than did the parental BCG strain. The isogenic,
      ***urease*** C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine,
     providing an intraphagosomal pH closer to the acidic pH optimum for Hly.
ΙT
of Organisms
       macrophage: immune system, blood and lymphatics
ΤТ
       tuberculosis: bacterial disease, drug therapy
        Tuberculosis (MeSH)
ΙT
    Chemicals & Biochemicals
           ***urease*** [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin:
        immunologic-drug, vaccine
RN
     9002-13-5 ( ***urease*** )
     9002-13-5 (EC 3.5.1.5)
    ANSWER 3 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
L2
     2004:927244 CAPLUS <<LOGINID::20080330>>
ΑN
    141:394066
DN
ΤI
    Vaccines comprising antigen domain and phagolysosomal escape domain for
     treating tuberculosis, cancer and infection
ΙN
       ***Grode, Leander*** ; Kaufmann, Stefan H. E.; Raupach, Baerbel; Hess,
     Juergen
    Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany
PA
SO
    PCT Int. Appl., 39 pp.
    CODEN: PIXXD2
DΤ
    Patent
    English
LA
FAN.CNT 1
                       KIND
                                          APPLICATION NO.
    PATENT NO.
                               DATE
                                                                DATE
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    WO 2004094469
                        A1
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            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
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    JP 2007524367
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	US	2007134267	A1	20070614	US	2006-554408	20061130
PRAI	US	2003-464644P	P	20030423			
	WO	2004-EP4345	W	20040423			

AB The present invention relates to novel recombinant vaccines comprising fusion protein contg. an antigenic domain and a phagolysosomal escape domain. providing protective immunity against tuberculosis. The antigenic domain is from Mycobacterium tuberculosis antigen Ag85B, Ag85A or ESAT-6; or Mycobacterium bovis antigen Ag85B. The antigenic domain can also be derived from autoantigen, tumor antigen, viral antigen, parasitic antigen, bacterial antigen or their immunogenic fragment. The phagolysosomal escape domain is a Listeria phagolysosomal escape domain. Further, the present invention refers to novel recombinant nucleic acid mols., vectors contg. said nucleic acid mols., cells transformed with said nucleic acid mols. and polypeptides encoded by said nucleic acid mols. These recombinant vaccines are used together with diluents, carriers and adjuvants; and are prepd. for mucosal or parenteral administration.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- IN ***Grode, Leander*** ; Kaufmann, Stefan H. E.; Raupach, Baerbel; Hess,
 Juergen
- IT 9002-13-5, ***Urease***

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(inactivation or -deficient; vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection)

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=> e kaufmann stefan h/au
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E1	1	KAUFMANN	STEFAN F M/AU	
E2	1	KAUFMANN	STEFAN G/AU	
E3	4>	KAUFMANN	STEFAN H/AU	
E4	965	KAUFMANN	STEFAN H E/AU	
E5	1	KAUFMANN	STEFAN H K/AU	
E6	4	KAUFMANN	STEFAN HE/AU	
E7	2	KAUFMANN	STEFAN HUGO ERNST/A	4U
E8	1	KAUFMANN	STEFAN J E/AU	
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E12	8	KAUFMANN	STEPHAN/AU	

=> s e3-e7 and (urease deficient)

L3 0 ("KAUFMANN STEFAN H"/AU OR "KAUFMANN STEFAN H E"/AU OR "KAUFMANN STEFAN HK"/AU OR "KAUFMANN STEFAN HUGO ERNST"/AU) AND (UREASE DEFICIENT)

=> s e3-e7 and (urease)

4 ("KAUFMANN STEFAN H"/AU OR "KAUFMANN STEFAN H E"/AU OR "KAUFMANN STEFAN HE"/AU OR "KAUFMANN STEFAN HUGO ERNST"/AU) AND (UREASE)

=> dup rem 14
PROCESSING COMPLETED FOR L4
L5 2 DUP REM L4 (2 DUPLICATES REMOVED)

=> d bib ab kwic 1-YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y/(N):y

- L5 ANSWER 1 OF 2 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>
- DN PREV200510303569
- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU Grode, Leander; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; ***Kaufmann, Stefan H. E.***
 [Reprint Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany Kaufmann@mpiib-Berlin.mpg.de
- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.

 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article
- LA English
- ED Entered STN: 23 Nov 2005 Last Updated on STN: 23 Nov 2005
- AΒ The tuberculosis vaccine Mycobacterium bovis bacille Calmette-Guerin (BCG) was equipped with the membrane-perforating listeriolysin (Hly) of Listeria monocytogenes, which was shown to improve protection against Mycobacterium tuberculosis. Following aerosol challenge, the Hly-secreting recombinant BCG (hly(+) rBCG) vaccine was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG ***urease*** C-deficient hly(+) rBCG (Delta strain. The isogenic, ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for Hly activity, exhibited still higher vaccine efficacy than parental BCG. Delta ureC hly(+) rBCG also induced profound protection against a member of the M. tuberculosis Beijing/W genotype family while parental BCG failed to do so consistently. Hly not only promoted antigen translocation into the cytoplasm but also apoptosis of infected macrophages. We concluded that superior vaccine efficacy of Delta ureC hly(+) rBCG as compared with parental BCG is primarily based on improved cross-priming, which causes enhanced T cell-mediated immunity.
- AU. . . Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; Raupach, Barbell; ***Kaufmann, Stefan H. E.*** [Reprint Author]
- AB. . . was shown to protect significantly better against aerosol infection with M. tuberculosis than did the parental BCG strain. The isogenic,

 urease C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the acidic pH optimum for Hly.
- IT . . .
- of Organisms

macrophage: immune system, blood and lymphatics

IT Diseases

tuberculosis: bacterial disease, drug therapy
Tuberculosis (MeSH)

IT Chemicals & Biochemicals

```
***urease*** [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin:
       immunologic-drug, vaccine
RN
     9002-13-5 ( ***urease*** )
     9002-13-5 (EC 3.5.1.5)
L5
    ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
ΑN
     2004:927244 CAPLUS <<LOGINID::20080330>>
DN
    141:394066
    Vaccines comprising antigen domain and phagolysosomal escape domain for
ΤТ
    treating tuberculosis, cancer and infection
ΙN
    Grode, Leander; ***Kaufmann, Stefan H. E.***; Raupach, Baerbel; Hess,
    Juergen
PΑ
    Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany
SO
    PCT Int. Appl., 39 pp.
    CODEN: PIXXD2
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    Patent
LA
    English
FAN.CNT 1
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     PATENT NO.
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    WO 2004094469
                               20041104 WO 2004-EP4345
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            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
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A1 20070614 US 2006-554408
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PRAI US 2003-464644P
                              20030423
     WO 2004-EP4345
                               20040423
    The present invention relates to novel recombinant vaccines comprising
AB
     fusion protein contq. an antigenic domain and a phagolysosomal escape
     domain. providing protective immunity against tuberculosis. The antigenic
     domain is from Mycobacterium tuberculosis antigen Ag85B, Ag85A or ESAT-6;
     or Mycobacterium bovis antigen Ag85B. The antigenic domain can also be
     derived from autoantigen, tumor antigen, viral antigen, parasitic antigen,
     bacterial antigen or their immunogenic fragment. The phagolysosomal
     escape domain is a Listeria phagolysosomal escape domain. Further, the
```

present invention refers to novel recombinant nucleic acid mols., vectors contq. said nucleic acid mols., cells transformed with said nucleic acid

mols. and polypeptides encoded by said nucleic acid mols. These recombinant vaccines are used together with diluents, carriers and

adjuvants; and are prepd. for mucosal or parenteral administration. RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- IN Grode, Leander; ***Kaufmann, Stefan H. E.*** ; Raupach, Baerbel; Hess,
 Juergen
- IT 9002-13-5, ***Urease***

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(inactivation or -deficient; vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection)

```
=> e raupach barbel/au
E1
           57
                RAUPACH B/AU
E2
           36
                 RAUPACH BAERBEL/AU
Е3
           22 --> RAUPACH BARBEL/AU
E4
           1
                 RAUPACH BARBELL/AU
E5
           7
                 RAUPACH C/AU
           7
Ε6
                 RAUPACH CARINA/AU
          8
                RAUPACH D C/AU
E7
E8
           5
                RAUPACH DALE C/AU
           1
                RAUPACH DALE R/AU
E9
           9
                RAUPACH E/AU
E10
         125
                RAUPACH F/AU
E11
E12
           2.
                RAUPACH F V/AU
```

=> s e1-e4 and urease

L6 4 ("RAUPACH B"/AU OR "RAUPACH BAERBEL"/AU OR "RAUPACH BARBEL"/AU
OR "RAUPACH BARBELL"/AU) AND UREASE

=> dup rem 16

PROCESSING COMPLETED FOR L6

L7 2 DUP REM L6 (2 DUPLICATES REMOVED)

=> d bib ab kwic 1-

YOU HAVE REQUESTED DATA FROM 2 ANSWERS - CONTINUE? Y/(N):y

- L7 ANSWER 1 OF 2 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN DUPLICATE 1 $\,$
- AN 2005:517633 BIOSIS <<LOGINID::20080330>>
- DN PREV200510303569
- TI Increased vaccine efficacy against tuberculosis of recombinant Mycobacterium bovis bacille Calmette-Guerin mutants that secrete listeriolysin.
- AU Grode, Leander; Seiler, Peter; Baumann, Sven; Hess, Juergen; Brinkmann, Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian; Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc; van Soolingen, Dick; ***Raupach, Barbell***; Kaufmann, Stefan H. E. [Reprint Author]
- CS Max Planck Inst Infect Biol, Dept Immunol, Schumannstr 21-22, D-10117 Berlin, Germany Kaufmann@mpiib-Berlin.mpg.de
- SO Journal of Clinical Investigation, (SEP 2005) Vol. 115, No. 9, pp. 2472-2479.

 CODEN: JCINAO. ISSN: 0021-9738.
- DT Article

```
LA
    English
    Entered STN: 23 Nov 2005
ED
    Last Updated on STN: 23 Nov 2005
    The tuberculosis vaccine Mycobacterium bovis bacille Calmette-Guerin (BCG)
AΒ
    was equipped with the membrane-perforating listeriolysin (Hly) of Listeria
    monocytogenes, which was shown to improve protection against Mycobacterium
     tuberculosis. Following aerosol challenge, the Hly-secreting recombinant
     BCG (hly(+) rBCG) vaccine was shown to protect significantly better
     against aerosol infection with M. tuberculosis than did the parental BCG
     strain. The isogenic, ***urease*** C-deficient hly(+) rBCG (Delta
     ureC hly(+) rBCG) vaccine, providing an intraphagosomal pH closer to the
     acidic pH optimum for Hly activity, exhibited still higher vaccine
     efficacy than parental BCG. Delta ureC hly(+) rBCG also induced profound
    protection against a member of the M. tuberculosis Beijing/W genotype
    family while parental BCG failed to do so consistently. Hly not only
    promoted antigen translocation into the cytoplasm but also apoptosis of
     infected macrophages. We concluded that superior vaccine efficacy of
     Delta ureC hly(+) rBCG as compared with parental BCG is primarily based on
     improved cross-priming, which causes enhanced T cell-mediated immunity.
    . . Volker; Eddine, Ali Nasser; Mann, Peggy; Goosmann, Christian;
    Bandermann, Silke; Smith, Debbie; Bancroft, Gregory J.; Reyrat, Jean-Marc;
    van Soolingen, Dick; ***Raupach, Barbell***; Kaufmann, Stefan H. E.
     [Reprint Author]
     . . was shown to protect significantly better against aerosol infection
    with M. tuberculosis than did the parental BCG strain. The isogenic,
      ***urease*** C-deficient hly(+) rBCG (Delta ureC hly(+) rBCG) vaccine,
    providing an intraphagosomal pH closer to the acidic pH optimum for Hly.
of Organisms
       macrophage: immune system, blood and lymphatics
ΙT
    Diseases
       tuberculosis: bacterial disease, drug therapy
       Tuberculosis (MeSH)
ΙT
    Chemicals & Biochemicals
           ***urease*** [EC 3.5.1.5]; listeriolysin; bacille Calmette-Guerin:
       immunologic-drug, vaccine
     9002-13-5 ( ***urease*** )
RN
    9002-13-5 (EC 3.5.1.5)
L7
    ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on STN
    2004:927244 CAPLUS <<LOGINID::20080330>>
ΑN
    141:394066
DN
ΤI
    Vaccines comprising antigen domain and phagolysosomal escape domain for
    treating tuberculosis, cancer and infection
ΙN
    Grode, Leander; Kaufmann, Stefan H. E.; ***Raupach, Baerbel*** ; Hess,
    Juergen
    Max-Planck-Gesellschaft zur Foerderung der Wissenschaften e.V., Germany
PA
    PCT Int. Appl., 39 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
    PATENT NO.
                       KIND DATE APPLICATION NO.
                                                               DATE
```

20041104 WO 2004-EP4345 20040423

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,

A1

WO 2004094469

PΤ

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CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
             SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
     AU 2004232485
                          Α1
                                20041104
                                            AU 2004-232485
                                                                    20040423
     CA 2523084
                          Α1
                                20041104
                                            CA 2004-2523084
                                                                    20040423
     EP 1618128
                          Α1
                                20060125
                                            EP 2004-729090
                                                                    20040423
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
     BR 2004009789
                                20060530
                                            BR 2004-9789
                          Α
                                                                    20040423
     CN 1798762
                                20060705
                                            CN 2004-80010664
                          Α
                                                                    20040423
     JP 2007524367
                          Τ
                                20070830
                                            JP 2006-505250
                                                                    20040423
                                            ZA 2005-8276
     ZA 2005008276
                          Α
                                20060628
                                                                    20051013
     IN 2005KN02337
                          Α
                                20070727
                                            IN 2005-KN2337
                                                                    20051122
                                            US 2006-554408
     US 2007134267
                          A1
                                20070614
                                                                    20061130
PRAI US 2003-464644P
                          Ρ
                                20030423
     WO 2004-EP4345
                         W
                                20040423
```

- AB The present invention relates to novel recombinant vaccines comprising fusion protein contg. an antigenic domain and a phagolysosomal escape domain. providing protective immunity against tuberculosis. The antigenic domain is from Mycobacterium tuberculosis antigen Ag85B, Ag85A or ESAT-6; or Mycobacterium bovis antigen Ag85B. The antigenic domain can also be derived from autoantigen, tumor antigen, viral antigen, parasitic antigen, bacterial antigen or their immunogenic fragment. The phagolysosomal escape domain is a Listeria phagolysosomal escape domain. Further, the present invention refers to novel recombinant nucleic acid mols., vectors contg. said nucleic acid mols., cells transformed with said nucleic acid mols. and polypeptides encoded by said nucleic acid mols. These recombinant vaccines are used together with diluents, carriers and adjuvants; and are prepd. for mucosal or parenteral administration.
- RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- IN Grode, Leander; Kaufmann, Stefan H. E.; ***Raupach, Baerbel*** ; Hess,
 Juergen
- IT 9002-13-5, ***Urease***

RL: BSU (Biological study, unclassified); REM (Removal or disposal); BIOL (Biological study); PROC (Process)

(inactivation or -deficient; vaccines comprising antigen domain and phagolysosomal escape domain for treating tuberculosis, cancer and infection)

```
=> e hess jurgen/au
E1
             1
                   HESS JUNIOR ARTUR/AU
             3
E2
                   HESS JURG/AU
Е3
            38 --> HESS JURGEN/AU
E4
             2
                   HESS JURGEN C/AU
E5
             2
                   HESS JURGEN H/AU
Ε6
             6
                   HESS JUSTIN M/AU
Ε7
           605
                   HESS K/AU
            23
                   HESS K A/AU
Ε8
```

```
1 HESS K BELLEVILLE F/AU
2 HESS K C/AU
E9
E10
E11
            8
                  HESS K D/AU
                HESS K G/AU
E12
            2
=> s e2-e5 and urease
            1 ("HESS JURG"/AU OR "HESS JURGEN"/AU OR "HESS JURGEN C"/AU OR
               "HESS JURGEN H"/AU) AND UREASE
=> d
L8
    ANSWER 1 OF 1 MEDLINE on STN
    2005580918
                 MEDLINE <<LOGINID::20080330>>
DN
    PubMed ID: 16110326
ΤI
    Increased vaccine efficacy against tuberculosis of recombinant
    Mycobacterium bovis bacille Calmette-Guerin mutants that secrete
     listeriolysin.
ΑU
    Grode Leander; Seiler Peter; Baumann Sven;
                                                ***Hess Jurgen*** ;
     Brinkmann Volker; Nasser Eddine Ali; Mann Peggy; Goosmann Christian;
     Bandermann Silke; Smith Debbie; Bancroft Gregory J; Reyrat Jean-Marc; van
     Soolingen Dick; Raupach Barbel; Kaufmann Stefan H E
    Max Planck Institute for Infection Biology, Berlin, Germany.
CS
    The Journal of clinical investigation, (2005 Sep) Vol. 115, No. 9, pp.
SO
     2472-9. Electronic Publication: 2005-08-18.
    Journal code: 7802877. ISSN: 0021-9738.
    United States
CY
    Journal; Article; (JOURNAL ARTICLE)
     (RESEARCH SUPPORT, NON-U.S. GOV'T)
LA
    English
FS
    Abridged Index Medicus Journals; Priority Journals
EΜ
    200512
ED
    Entered STN: 3 Nov 2005
    Last Updated on STN: 18 Dec 2005
     Entered Medline: 14 Dec 2005
=> s (urease deficient)
           75 (UREASE DEFICIENT)
=> dup rem 19
PROCESSING COMPLETED FOR L9
            31 DUP REM L9 (44 DUPLICATES REMOVED)
=> s 110 and (bact? or mycobact? or tuberculosis or bovis)
            20 L10 AND (BACT? OR MYCOBACT? OR TUBERCULOSIS OR BOVIS)
=> d bib ab kwic 1-
YOU HAVE REQUESTED DATA FROM 20 ANSWERS - CONTINUE? Y/(N):y
L11 ANSWER 1 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
     2007:355406 BIOSIS <<LOGINID::20080330>>
AN
    PREV200700359871
TΙ
    Characterization of the urease operon of Brucella abortus and assessment
     of its role in virulence of the ***bacterium***
ΑU
    Sangari, Felix J.; Seoane, Asuncion; Rodriguez, Maria Cruz; Aguero, Jesus;
    Garcia Lobo, Juan M. [Reprint Author]
CS
    Univ Cantabria, Dept Biol Mol, Fac Med, C Cardenal Herrera Oria S-N,
```

Santander 39011, Spain jmglobo@unican.es

- SO Infection and Immunity, (FEB 2007) Vol. 75, No. 2, pp. 774-780. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 20 Jun 2007 Last Updated on STN: 20 Jun 2007
- AΒ Most members of the genus Brucella show strong urease activity. However, the role of this enzyme in the pathogenesis of Brucella infections is poorly understood. We isolated several Tn5 insertion mutants deficient in urease activity from Brucella abortus strain 2308. The mutations of most of these mutants mapped to a 5.7-kbp DNA region essential for urease activity. Sequencing of this region, designated urel, revealed the presence of seven open reading frames corresponding to the urease structural proteins (UreA, UreB, and UreC) and the accessory proteins (UreD, UreE, UreF, and UreG). In addition to the urease genes, another gene (cobT) was identified, and inactivation of this gene affected urease activity in Brucella. Subsequent analysis of the previously described sequences of the genomes of Brucella spp. revealed the presence of a second urease cluster, ure2, in all them. The ure2 locus was apparently ***Urease*** - ***deficient*** mutants inactive in B. abortus 2308. were used to evaluate the role of urease in Brucella pathogenesis. The urease-producing strains were found to be resistant in vitro to strong acid conditions in the presence of urea, while urease-negative mutants were susceptible to acid treatment. Similarly, the urease-negative mutants were killed more efficiently than the urease-producing strains during transit through the stomach. These results suggested that urease protects brucellae during their passage through the stomach when the ***bacteria*** are acquired by the oral route, which is the major route of infection in human brucellosis.
- TI Characterization of the urease operon of Brucella abortus and assessment of its role in virulence of the $\,$ ***bacterium*** .
- AB. . . presence of a second urease cluster, ure2, in all them. The ure2 locus was apparently inactive in B. abortus 2308. ***Urease*** ***deficient*** mutants were used to evaluate the role of urease in Brucella pathogenesis. The urease-producing strains were found to be resistant. . . during transit through the stomach. These results suggested that urease protects brucellae during their passage through the stomach when the ***bacteria*** are acquired by the oral route, which is the major route of infection in human brucellosis.
- IT . . .

and Assimilation); Enzymology (Biochemistry and Molecular Biophysics)

IT Parts, Structures, & Systems of Organisms

stomach: digestive system

IT Diseases

brucellosis: ***bacterial*** disease, infectious disease
Brucellosis (MeSH)

IT Diseases

Brucella abortus infection: ***bacterial*** disease, infectious disease

IT Chemicals & Biochemicals

DNA; urease [EC 3.5.1.5]; UreA; UreB; UreG; UreD; UreE; UreF; UreC ORGN Classifier

Gram-Negative Aerobic Rods and Cocci 06500

Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

Organism Name
Brucella abortus (species): strain-2308
Taxa Notes
Bacteria , Eubacteria, Microorganisms
ORGN Classifier
Hominidae 86215
Super Taxa
Primates; Mammalia; Vertebrata; Chordata; Animalia
Organism Name
human (common)
Taxa Notes
Animals, . . .

- L11 ANSWER 2 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2006:296770 BIOSIS <<LOGINID::20080330>>
- DN PREV200600297562
- TI The role of Klebsiella pneumoniae urease in intestinal colonization and resistance to gastrointestinal stress.
- AU Maroncle, Nathalie; Rich, Chantal; Forestier, Christiane [Reprint Author]
- CS Univ Auvergne, Fac Pharm, Bacteriol Lab, 28 Pl H Dunant, F-63000 Clermont Ferrand, France
 - Christiane.forestier@u-clermontI.fr
- SO Research in Microbiology, (MAR 2006) Vol. 157, No. 2, pp. 184-193. CODEN: RMCREW. ISSN: 0923-2508.
- DT Article
- LA English
- ED Entered STN: 31 May 2006 Last Updated on STN: 31 May 2006
- AB The first step in nosocomial infections due to Klebsiella pneumoniae is colonization of the patient's gastrointestinal (GI) tract. In a previous work, signature-tagged mutagenesis was used in a murine model to identify 13 genes required for efficient colonization, two of which were involved in urea metabolism. The role of urease was further investigated by the construction and analysis of an isogenic ***urease*** -
 - ***deficient*** mutant. The behavior of both the wild-type strain and ***urease*** - ***deficient*** mutant was tested under hostile conditions, reproducing stresses encountered in the GI environment. The wild-type strain had an acid tolerance response (ATR) to inorganic acid, was resistant to organic acids (38.5% survival) and was able to survive concentrations of bile encountered in vivo. The absence of urease did not affect the resistance of K. pneumoniae to acid and bile stresses, but the enhanced adhesion response to Int-407 cells after exposure to bile observed with the wild-type strain was no longer detected with the urease mutant. When tested in the murine intestinal colonization model, both strains were mainly recovered in the large intestine parts, and the mutant was impaired in its colonization capacities, but only when tested in competition with the wild-type strain. These findings emphasize the prominent role played by metabolic function in the colonization process of such a complex ecosystem as the host GI tract. (c) 2005 Elsevier SAS. All rights reserved.
- AB. . . were involved in urea metabolism. The role of urease was further investigated by the construction and analysis of an isogenic
 urease ***deficient*** mutant. The behavior of both the
 wild-type strain and the ***urease*** ***deficient*** mutant was
 tested under hostile conditions, reproducing stresses encountered in the
 GI environment. The wild-type strain had an acid tolerance. . .

 ORGN Classifier

Enterobacteriaceae 06702 Super Taxa

Facultatively Anaerobic Gram-Negative Rods; Eubacteria;
 Bacteria ; Microorganisms

Organism Name

Klebsiella pneumoniae (species): pathogen

Taxa Notes

Bacteria , Eubacteria, Microorganisms

ORGN Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

human (common): host

Taxa Notes

- L11 ANSWER 3 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2006:26018 BIOSIS <<LOGINID::20080330>>
- DN PREV200600025071
- TI Production of ammonium by Helicobacter pylori mediates occludin processing and disruption of tight junctions in Caco-2 cells.
- AU Lytton, Simon D. [Reprint Author]; Fischer, Wolfgang; Nagel, Wolfram; Haas, Rainer; Beck, Franz X.
- CS SeraDiaLogist, Hertlingstr 1, D-81545 Munich, Germany Simon.lytton@t-online.de
- SO Microbiology (Reading), (OCT 2005) Vol. 151, No. Part 10, pp. 3267-3276. ISSN: 1350-0872.
- DT Article
- LA English
- ED Entered STN: 21 Dec 2005 Last Updated on STN: 21 Dec 2005
- Tight junctions, paracellular permeability barriers that define epithelial AΒ cell polarity, play an essential role in transepithelial transport, cell-cell adhesion and lymphocyte transmigration. They are also important for the maintenance of innate immune defence and intestinal antigen uptake. Ammonium (NH4+) is elevated in the gastric aspirates of Helicobacter pylori-infected patients and has been implicated in the disruption of tight-junction functional integrity and the induction of gastric mucosal damage during H. pylori infection. The precise mechanism of the effect of ammonium and the molecular targets of ammonium in host tissue are not yet identified. To study the effects of ammonium on epithelial tight junctions, the human colon carcinoma cell line Caco-2 was cultured on permeable supports and the transepithelial resistance (TER) was measured at different time intervals following exposure to ammonium salts or H. pylori-derived ammonium. A biphasic response to treatment with ammonium was found. Acute exposure to ammonium salts or NH3/NH4+ derived from urea metabolism by wild-type H. pylori resulted in a 20-30% decrease in TER. After 24 h, the NH4Cl-treated cells showed a partial recovery of TER. In contrast, the control culture, or cultures that were exposed to supernatants derived from ***urease*** - ***deficient*** H. pylori, showed no significant decrease in TER. Occludin-specific immunoblots revealed the expression of a low-molecular-weight form of occludin of 42 kDa upon NH3/NH4+ exposure. The results indicate that modulation of tight-junction function by H. pylori is ammonium-dependent and linked to the accumulation of a low-molecular-weight and detergent-soluble form of occludin.
- AB. . . showed a partial recovery of TER. In contrast, the control culture,

```
or cultures that were exposed to supernatants derived from ***urease***
     - ***deficient*** H. pylori, showed no significant decrease in TER.
    Occludin-specific immunoblots revealed the expression of a
     low-molecular-weight form of occludin of.
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
     Super Taxa
       Eubacteria; ***Bacteria***; Microorganisms
     Organism Name
       Helicobacter pylori (species): pathogen
     Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Hominidae
                   86215
     Super Taxa
       Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
       human (common)
       Caco-2 cell line (cell_line). . .
L11 ANSWER 4 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    2004:316567 BIOSIS <<LOGINID::20080330>>
ΑN
    PREV200400316839
DN
    Selection and properties of Streptococcus thermophilus mutants deficient
ΤI
    in urease.
ΑIJ
    Monnet, C. [Reprint Author]; Pernoud, S.; Sepulchre, A.; Fremaux, C.;
    Corrieu, G.
CS
    Unite Mixte Rech Genie and Microbiol Proc Alimentai, INRA, F-78850,
    Thiverval Grignon, France
     monnet@grignon.inra.fr
    Journal of Dairy Science, (June 2004) Vol. 87, No. 6, pp. 1634-1640.
    print.
    CODEN: JDSCAE. ISSN: 0022-0302.
DT
    Article
LA
    English
ED
    Entered STN: 15 Jul 2004
    Last Updated on STN: 15 Jul 2004
    Natural variations of the urea content of milk have a detrimental effect
AΒ
     on the regularity of acidification by Streptococcus thermophilus strains
     used in dairy processes. The aim of the present study was to select
       ***urease*** - ***deficient*** mutants of S. thermophilus and to
     investigate their properties. Using an improved screening medium on agar
     plates, mutants were selected from 4 different parent strains after
    mutagen treatment and by spontaneous mutation. Most mutants were stable
     and had a phage sensitivity profile similar to that of their parent
     strain. Some of them contained detrimental secondary mutations, as their
     acidifying activity was lower than that of the parent strain cultivated in
     the presence of the urease inhibitor flurofamide. The proportion of this
     type of mutant was much lower among spontaneous mutants than among mutants
     selected after mutagen treatment. Utilization of ***urease*** -
      ***deficient*** mutants in dairy processes may have several advantages,
     such as an increase in acidification, an improved regularity of
     acidification, and a lower production of ammonia in whey.
AB.
    . . regularity of acidification by Streptococcus thermophilus strains
    used in dairy processes. The aim of the present study was to select
       ***urease*** - ***deficient*** mutants of S. thermophilus and to
```

investigate their properties. Using an improved screening medium on agar

```
plates, mutants were selected. . . of this type of mutant was much
    lower among spontaneous mutants than among mutants selected after mutagen
    treatment. Utilization of ***urease*** - ***deficient*** mutants in
    dairy processes may have several advantages, such as an increase in
     acidification, an improved regularity of acidification, and. . .
ORGN Classifier
       Gram-Positive Cocci
                             07700
     Super Taxa
                    ***Bacteria*** ; Microorganisms
       Eubacteria;
     Organism Name
       Streptococcus thermophilus (species): ***urease*** -
          ***deficient*** mutants
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
    ANSWER 5 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
L11
AN
     2004:104625 BIOSIS <<LOGINID::20080330>>
DN
    PREV200400096230
ΤI
                  ***urease*** - ***deficient*** derivatives of
    Motility of
    Helicobacter pylori.
ΑU
    Tan, Shumin; Berg, Douglas E. [Reprint Author]
CS
    Department of Molecular Microbiology, Washington University School of
    Medicine, Campus Box 8230, St. Louis, MO, 63110, USA
    berg@borcim.wustl.edu
SO
    Journal of Bacteriology, (February 2004) Vol. 186, No. 3, pp. 885-888.
    CODEN: JOBAAY. ISSN: 0021-9193.
DT
    Article
LA
    English
    Entered STN: 18 Feb 2004
ED
    Last Updated on STN: 18 Feb 2004
    Early studies of a ureB mutant derivative of Helicobacter pylori had
AB
     suggested that urease is needed for motility and that urease action helps
     energize flagellar rotation. Here we report experiments showing that
    motility is unaffected by deletion of ureA and ureB (urease genes) or by
     inactivation of ureB alone, especially if H. pylori strains used as
     recipients for transformation with mutant alleles are preselected for
    motility. This result was obtained with the strain used in the early
     studies (CPY3401) and also with 15 other strains, 3 of which can colonize
    mice. We conclude that urease is not needed for H. pylori motility.
    Motility of ***urease*** - ***deficient*** derivatives of
    Helicobacter pylori.
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives
     Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
     Organism Name
       Helicobacter pylori (species): pathogen, motility, strain-88-3887,
       strain-A28-1, strain-A66-1, strain-CYP3401, strain-Chen13, strain-F28,
       strain-GS5, strain-HK192, strain-PCM4, strain-PeCan28, strain-R64,
       strain-R66, strain-R76, strain-R82, strain-SS1, strain-X47,
          ***urease*** - ***deficient*** derivatives
     Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Muridae 86375
     Super Taxa
```

Rodentia; Mammalia; Vertebrata; Chordata; Animalia Organism Name mouse (common) Taxa Notes Animals,. . .

- L11 ANSWER 6 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2000:418052 BIOSIS <<LOGINID::20080330>>
- DN PREV200000418052
- TI Dual functions of Streptococcus salivarius urease.
- AU Chen, Yi-Ywan M.; Weaver, Cheryl A.; Burne, Robert A. [Reprint author]
- CS Center for Oral Biology, University of Rochester Medical Center, 601 Elmwood Ave., Rochester, NY, 14642, USA
- SO Journal of Bacteriology, (August, 2000) Vol. 182, No. 16, pp. 4667-4669. print.

 CODEN: JOBAAY. ISSN: 0021-9193.
- DT Article
- LA English
- ED Entered STN: 4 Oct 2000 Last Updated on STN: 8 Jan 2002
- AB A ***urease*** ***deficient*** derivative of Streptococcus salivarius 57.I was constructed by allelic exchange at the ureC locus. The wild-type strain was protected against acid killing through hydrolysis of physiologically relevant concentrations of urea, whereas the mutant was not. Also, S. salivarius could use urea as a source of nitrogen for growth exclusively through a urease-dependent pathway.
- AB A ***urease*** ***deficient*** derivative of Streptococcus salivarius 57.I was constructed by allelic exchange at the ureC locus. The wild-type strain was protected against. . .
- ORGN Classifier

Gram-Positive Cocci 07700

Super Taxa

Eubacteria; ***Bacteria***; Microorganisms

Organism Name

Streptococcus salivarius: strain-57.I

Taxa Notes

Bacteria , Eubacteria, Microorganisms

- L11 ANSWER 7 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 2000:400579 BIOSIS <<LOGINID::20080330>>
- DN PREV200000400579
- TI Helicobacter pylori urease suppresses ***bactericidal*** activity of peroxynitrite via carbon dioxide production.
- AU Kuwahara, Hideo; Miyamoto, Yoichi; Akaike, Takaaki [Reprint author]; Kubota, Tatsuo; Sawa, Tomohiro; Okamoto, Shinichiro; Maeda, Hiroshi [Reprint author]
- CS Department of Microbiology, Kumamoto University School of Medicine, 2-2-1 Honjo, Kumamoto, 860-0811, Japan
- SO Infection and Immunity, (August, 2000) Vol. 68, No. 8, pp. 4378-4383. print.

 CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 20 Sep 2000 Last Updated on STN: 8 Jan 2002
- AB Helicobacter pylori can produce a persistent infection in the human stomach, where chronic and active inflammation, including the infiltration

of phagocytes such as neutrophils and monocytes, is induced. H. pylori may have a defense system against the antimicrobial actions of phagocytes. We studied the defense mechanism of H. pylori against host-derived ***bactericidal*** metabolite of nitric peroxynitrite (ONOO-), a oxide, focusing on the role of H. pylori urease, which produces CO2 and NH3 from urea and is known to be an essential factor for colonization. The viability of H. pylori decreased in a time-dependent manner with continuous exposure to 1 muM ONOO-, i.e., 0.2% of the initial ***bacteria*** remained after a 5-min treatment without urea. The ***bactericidal*** action of ONOO- against H. pylori was significantly attenuated by the addition of 10 mM urea, the substrate for urease, whereas ONOO--induced killing of a ***urease*** - ***deficient*** mutant of H. pylori or Campylobacter jejuni, another microaerophilic ***bacterium*** lacking urease, was not affected by the addition of urea. Such as protective effect of urea was potentiated by supplementation with exogenous urease, and it was almost completely nullified by 10 muM flurofamide, a specific inhibitor of urease. The ***bactericidal*** action of ONOO- was also suppressed by the addition of 20 mM NaHCO3 but not by the addition of 20 mM NH3. In addition, the nitration of L-tyrosine of H. pylori after treatment with ONOO- was significantly reduced by the addition of urea or NaHCO3, as assessed by high-performance liquid chromatography with electrochemical detection. These results suggest that H. pylori-associated urease functions to produce a potent ONOO- scavenger, CO2/HCO3-, that defends the ***bacteria*** from ONOO- cytotoxicity. The protective effect of urease

may thus facilitate sustained ***bacterial*** colonization in the
infected gastric mucosa.

- TI Helicobacter pylori urease suppresses ***bactericidal*** activity of peroxynitrite via carbon dioxide production.
- . . system against the antimicrobial actions of phagocytes. We studied the defense mechanism of H. pylori against host-derived peroxynitrite ***bactericidal*** metabolite of nitric oxide, focusing on (ONOO-), a the role of H. pylori urease, which produces CO2 and NH3 from urea and. . of H. pylori decreased in a time-dependent manner with continuous exposure to 1 muM ONOO-, i.e., 0.2% of the initial ***bacteria*** remained after a 5-min treatment without urea. The ***bactericidal*** action of ONOO- against H. pylori was significantly attenuated by the addition of 10 mM urea, the substrate for urease, whereas ONOO--induced ***urease*** - ***deficient*** mutant of H. pylori or killing of a Campylobacter jejuni, another microaerophilic ***bacterium*** urease, was not affected by the addition of urea. Such as protective effect of urea was potentiated by supplementation with exogenous urease, and it was almost completely nullified by 10 muM flurofamide, a specific inhibitor of urease. The ***bactericidal*** action of ONOO- was also suppressed by the addition of 20 mM NaHCO3 but not by the addition of 20. . . electrochemical detection. These results suggest that H. pylori-associated urease functions to produce a potent ONOO- scavenger, CO2/HCO3-, that defends the ***bacteria*** from ONOO- cytotoxicity. The protective effect of urease may thus facilitate sustained ***bacterial*** colonization in the infected gastric mucosa.

IT . . Organisms

gastric mucosa: digestive system, infection; phagocytes: immune system; stomach: digestive system

IT Chemicals & Biochemicals

carbon dioxide: production; peroxynitrite: ***bactericidal***

```
activity, nitric oxide ***bactericidal*** metabolite; urease:
       Helicobacter pylori
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
     Super Taxa
       Eubacteria; ***Bacteria***; Microorganisms
     Organism Name
       Campylobacter jejuni: pathogen
       Helicobacter pylori: defense mechanism, pathogen
     Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Hominidae 86215
     Super Taxa
       Primates; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
       human
     Taxa Notes
       Animals, Chordates, . . .
L11 ANSWER 8 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
    1999:114559 BIOSIS <<LOGINID::20080330>>
ΑN
    PREV199900114559
DN
    Genetic and physiologic characterization of urease of Actinomyces
ΤI
    naeslundii.
ΑIJ
    Morou-Bermudez, Evangelia; Burne, Robert A. [Reprint author]
CS
    Cent. Oral Biol., Univ. Rochester Med. Cent., 601 Elmwood Ave., Rochester,
    NY 14642, USA
    Infection and Immunity, (Feb., 1999) Vol. 67, No. 2, pp. 504-512. print.
SO
    CODEN: INFIBR. ISSN: 0019-9567.
DT
    Article
LA
    English
    Entered STN: 12 Mar 1999
ED
    Last Updated on STN: 12 Mar 1999
AΒ
    Ammonia production from urea by ureolytic oral ***bacteria***
    believed to have a significant impact on oral health and the ecological
     balance of oral microbial populations. In this study we cloned and
     characterized the urease gene cluster of Actinomyces naeslundii, which is
     one of the pioneer organisms in the oral cavity and a significant
     constituent of supragingival and subgingival dental plaque in children and
     adults. An internal fragment of the ureC gene of A. naeslundii WVU45 was
     initially amplified by PCR with degenerate primers derived from conserved
     amino acid sequences of the large catalytic subunit of urease in
       ***bacteria*** and plants. The PCR product was then used as a probe to
     identify recombinant
                           ***bacteriophages*** carrying the A. naeslundii
     urease gene cluster and roughly 30 kbp of flanking DNA. Nucleotide
     sequence analysis demonstrated that the gene cluster was comprised of
     seven contiguously arranged open reading frames with significant
     homologies at the protein and nucleotide sequence levels to the ureABCEFGD
     genes from other organisms. By using primer extension, a putative
     transcription initiation site was mapped at 66 bases 5' to the start codon
     of ureA. A ***urease*** - ***deficient*** strain was constructed by
     insertion of a kanamycin resistance determinant within the ureC gene via
     allelic replacement. In contrast to the wild-type organism, the isogenic
    mutant was unable to grow in a semidefined medium supplemented with urea
    as the nitrogen source and was not protected by the addition of urea
     against killing in moderately acidic environments. These data indicated
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that urea can be effectively utilized as a nitrogen source by A. naeslundii via a urease-dependent pathway and that ureolysis can protect A. naeslundii against environmental acidification at physiologically relevant pH values. Therefore, urease could confer to A. naeslundii critical selective advantages over nonureolytic organisms in dental plaque, constituting an important determinant of plaque ecology. ***bacteria*** Ammonia production from urea by ureolytic oral believed to have a significant impact on oral health and the ecological balance of oral microbial populations. In this. . . amplified by PCR with degenerate primers derived from conserved amino acid sequences of the large catalytic subunit of urease in ***bacteria*** and plants. The PCR product was then used as a probe to identify recombinant ***bacteriophages*** carrying the A. naeslundii urease gene cluster and roughly 30 kbp of flanking DNA. Nucleotide sequence analysis demonstrated that the. . . primer extension, a putative transcription initiation site was mapped at 66 bases 5' to the start codon of ureA. A ***urease*** - ***deficient*** strain was constructed by insertion of a kanamycin resistance determinant within the ureC gene via allelic replacement. In contrast to. . ITMajor Concepts Enzymology (Biochemistry and Molecular Biophysics); Infection ΙT Diseases dental plaque: ***bacterial*** disease, dental and oral disease Dental Plaque (MeSH) Chemicals & Biochemicals ΤT ammonia: production; urea; urease; Actinomyces naeslundii ureA gene; Actinomyces. ORGN Classifier Irregular Nonsporing Gram-Positive Rods 08890 Actinomycetes and Related Organisms; Eubacteria; ***Bacteria*** ; Microorganisms Organism Name Actinomyces naeslundii: pathogen, strain-WVU45 Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms L11 ANSWER 9 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN ΑN 1996:361212 BIOSIS <<LOGINID::20080330>> DN PREV199699083568 Factors affecting growth and antibiotic susceptibility of Helicobacter pylori: Effect of pH and urea on the survival of a wild-type strain and a ***urease*** - ***deficient*** mutant. Sjostrom, J. E. [Reprint author]; Larsson, H. ΑU CS Dep. Cell Biol., Astra Hassle AB, Molndal, Sweden SO Journal of Medical Microbiology, (1996) Vol. 44, No. 6, pp. 425-433. CODEN: JMMIAV. ISSN: 0022-2615. DT Article LA English Entered STN: 14 Aug 1996 EDLast Updated on STN: 15 Aug 1996 AΒ This study investigated how pH and the presence of urea affect the survival and growth of Helicobacter pylori and whether these factors affect susceptibility to antibiotics in vitro. The viability of a wild-type strain and a ***urease*** - ***deficient*** mutant of H. pylori was studied after incubation for 1 h in buffers at different pH values at 37 degree C under microaerophilic conditions. Viable counts

were not affected at pH 5 and pH 7. In buffer at pH 3, there were no viable organisms, but urea (6.25 mm) protected the wild-type strain, which survived well. At pH 9, urea further reduced the viability of H. pylori and flurofamide almost abolished the effect of urea on the wild-type strain. Neither urea nor flurofamide affected the viability of the ***urease*** - ***deficient*** mutant under the same conditions. Growth was also pH dependent and was enhanced in shake-cultures. At pH 5, urea supported growth of the wild-type strain, but at pH 7 a toxic effect on the ***bacteria*** was observed. Growth of H. pylori at pH 5.9 was poor, and susceptibility to amoxycillin, erythromycin and clarithromycin was markedly less than at pH 7.2 and 7.9. The ***bactericidal*** activities of metronidazole and tetracycline were similar at the different pH values studied. At neutral pH the killing rates of amoxycillin and clarithromycin were growth rate dependent. Susceptibility to metronidazole was enhanced in stationary cultures. The interaction obtained between the proton pump inhibitor, omeprazole, and amoxycillin at pH 7 was of additive type. These results suggest that pH and growth conditions may be important in the antibacterial efficacy of different antibiotics in vivo and also provide a possible explanation for the potentiating effect of omeprazole with antibiotics in the treatment of H. pylori infections.

- TI. . . and antibiotic susceptibility of Helicobacter pylori: Effect of pH and urea on the survival of a wild-type strain and a ***urease*** ***deficient*** mutant.
- AB. . . Helicobacter pylori and whether these factors affect susceptibility to antibiotics in vitro. The viability of a wild-type strain and a ***urease*** ***deficient*** mutant of H. pylori was studied after incubation for 1 h in buffers at different pH values at 37 degree. . . flurofamide almost abolished the effect of urea on the wild-type strain. Neither urea nor flurofamide affected the viability of the ***urease*** ***deficient*** mutant under the same conditions. Growth was also pH dependent and was enhanced in shake-cultures. At pH 5, urea supported growth of the wild-type strain, but at pH 7 a toxic effect on the ***bacteria*** was observed. Growth of H. pylori at pH 5.9 was poor, and susceptibility to amoxycillin, erythromycin and clarithromycin was markedly less than at pH 7.2 and 7.9. The ***bactericidal*** activities of metronidazole and tetracycline were similar at the different pH values studied. At neutral pH the killing rates of. . .

ORGN Classifier

Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

Organism Name

aerobic helical or vibrioid gram-negative ***bacteria***
Helicobacter pylori

Taxa Notes

Bacteria , Eubacteria, Microorganisms

ORGN Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

human

Taxa Notes

Animals, Chordates, . . .

L11 ANSWER 10 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on

STN

- AN 1996:183710 BIOSIS <<LOGINID::20080330>>
- DN PREV199698739839
- TI In vitro antibacterial activity of omeprazole and its selectivity for Helicobacter spp. are dependent on incubation conditions.
- AU Sjostrom, J. E. [Reprint author]; Fryklund, J.; Kuhler, T.; Larsson, H.
- CS Astra Hassle AB, Dep. Cell Biol., S-431 83 Molndal, Sweden
- SO Antimicrobial Agents and Chemotherapy, (1996) Vol. 40, No. 3, pp. 621-626. CODEN: AMACCQ. ISSN: 0066-4804.
- DT Article
- LA English
- ED Entered STN: 29 Apr 1996 Last Updated on STN: 29 Apr 1996
- AΒ Factors affecting the in vitro antibacterial activity of omeprazole were studied. Our data show that 3H-labeled omeprazole covalently bound to Helicobacter pylori and to other gram-negative and gram-positive ***bacteria*** . The compound was found to bind to a broad range of proteins in H. pylori, and at pH 5, binding was enhanced 15-fold compared with binding at pH 7. The ***bactericidal*** activity correlated to the degree of binding, and at pH 5, a pH at which omeprazole readily converts to the active sulfenamide form, beta-mercaptoethanol, a known scavenger of sulfenamide, and fetal calf serum, to which noncovalent protein binding of omeprazole is known to occur, reduced the level of binding and almost entirely abolished the ***bactericidal*** activity. At pH 7 the killing activities of omeprazole and structural analogs (e.g., proton pump inhibitors) were dependent on the time-dependent conversion (half-life) to the corresponding sulfenamide. The ***bactericidal*** activity exerted by the sulfenamide form at pH 5 was not specific for the genus Helicobacter. However, in brucella broth at pH 7 with 10% fetal calf serum, only Helicobacter spp. were susceptible to omeprazole, with MBCs in the range of 32 to 64 mu-g/ml, and MBCs for more stable proton pump inhibitors were higher. Wild-type H. pylori and its isogenic ***urease*** - ***deficient*** mutant were equally susceptible to omeprazole. Thus, the urease is not a lethal target for omeprazole action in H. pylori. In conclusion, the antibacterial activities of omeprazole and analogs are dependent on pH and the composition of the medium used. Thus, at a low pH in buffer, these compounds have a nonselective action, whereas in broth at neutral pH, the mechanism of action is selective for Helicobacter spp.
- AB. . . omeprazole were studied. Our data show that 3H-labeled omeprazole covalently bound to Helicobacter pylori and to other gram-negative and ***bacteria*** . The compound was found to bind to a gram-positive broad range of proteins in H. pylori, and at pH 5, binding was enhanced 15-fold compared with binding at pH 7. The ***bactericidal*** activity correlated to the degree of binding, and at pH 5, a pH at which omeprazole readily converts to the. . . which noncovalent protein binding of omeprazole is known to occur, reduced the level of binding and almost entirely abolished the ***bactericidal*** activity. At pH 7 the killing activities of omeprazole and structural analogs (e.g., proton pump inhibitors) were dependent on the time-dependent conversion (half-life) to the corresponding sulfenamide. The ***bactericidal*** activity exerted by the sulfenamide form at pH 5 was not specific for the genus Helicobacter. However, in brucella broth. . . 32 to 64 mu-q/ml, and MBCs for more stable proton pump inhibitors were higher. Wild-type H. pylori and its isogenic ***urease*** - ***deficient*** mutant were equally susceptible to omeprazole. Thus, the urease is not a lethal target for omeprazole action in H. pylori.. . .

```
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
     Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
     Organism Name
       aerobic helical or vibrioid gram-negative ***bacteria***
       Helicobacter pylori
     Taxa Notes
           ***Bacteria*** , Eubacteria, Microorganisms
L11 ANSWER 11 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
ΑN
    1995:315763 BIOSIS <<LOGINID::20080330>>
DN
    PREV199598330063
    Avirulent, ***urease*** - ***deficient*** Helicobacter pylori
TΤ
    colonizes gastric epithelial explants ex vivo.
ΑU
    Eaton, K. A. [Reprint author]; Krakowka, S.
CS
    Dep. Vet. Pathobiol., OSU, 1925 Coffey Rd., Columbus, OH 43210, USA
SO
    Scandinavian Journal of Gastroenterology, (1995) Vol. 30, No. 5, pp.
     434-437.
    CODEN: SJGRA4. ISSN: 0036-5521.
DT
    Article
LA
    Enalish
    Entered STN: 30 Jul 1995
ED
    Last Updated on STN: 30 Jul 1995
AB
    Background: Urease-negative Helicobacter pylori generated by insertional
    mutagenesis fails to colonize gnotobiotic piglets, and this effect is
    largely independent of gastric pH. The purpose of this study was to
    determine whether urease-negative H. pylori colonizes gastric explants ex
    vivo. Methods: Gastric mucosal explants derived from neonatal germ-free
    piglets were inoculated with either wild-type H. pylori or one of two
    mutants derived by insertional mutagenesis. Results: All three
      ***bacterial***
                       strains colonized explants. The level of colonization
     increased over the duration of the experiment, reaching 10-8-10-9 cfu/g
    gastric mucosa by 72 h after inoculation. Morphologic evidence of
     colonization was similar to that observed in gnotobiotic piglets.
    Conclusions: Colonization of explants was not affected by lack of urease.
    These results contrast with previous findings showing that urease activity
    is essential for colonization of piglets by H. pylori. Thus,
    urease-dependent colonization is dependent on an intact gastric
    microenvironment.
                 ***urease*** - ***deficient***
                                                  Helicobacter pylori
TΤ
    Avirulent,
     colonizes gastric epithelial explants ex vivo.
    . . piglets were inoculated with either wild-type H. pylori or one of
    two mutants derived by insertional mutagenesis. Results: All three
       ***bacterial*** strains colonized explants. The level of colonization
     increased over the duration of the experiment, reaching 10-8-10-9 cfu/g
    gastric mucosa by. .
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives 06210
     Super Taxa
       Eubacteria; ***Bacteria*** ; Microorganisms
     Organism Name
       aerobic helical or vibrioid gram-negative ***bacteria***
       Helicobacter pylori
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
```

ORGN Classifier
Suidae 85740
Super Taxa
Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia
Organism Name
pig
Taxa Notes
Animals, Artiodactyls, . . .

- L11 ANSWER 12 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on STN
- AN 1994:446969 BIOSIS <<LOGINID::20080330>>
- DN PREV199497459969
- TI Effect of gastric pH on urease-dependent colonization of gnotobiotic piglets by Helicobacter pylori.
- AU Eaton, Kathryn A. [Reprint author]; Krakowka, Steven
- CS Dep. Veterinary Pathobiol., Ohio State Univ., 1925 Coffey Road, Columbus, OH 43210, USA
- SO Infection and Immunity, (1994) Vol. 62, No. 9, pp. 3604-3607. CODEN: INFIBR. ISSN: 0019-9567.
- DT Article
- LA English
- ED Entered STN: 24 Oct 1994
 Last Updated on STN: 25 Oct 1994
- Thirty-seven gnotobiotic piglets from seven litters were infected with AB either Helicobacter pylori N6 or urease-negative H. pylori N6ureG::Km, which contains an insertion in the ureG gene and produces inactive urease. To produce achlorhydria, piglets were treated throughout the experiment with omeprazole (5 mg intravenously every 12 h) and ranitidine (75 mg orally every 6 h). Treatment resulted in elevation of gastric pH to 7.0 +- 1.1 throughout the experiment. Control piglets were not treated and remained normochlorhydric. Strain N6 colonized well in both normal and achlorhydric piglets. All 10 piglets were colonized, and colonization ranged from 4.4 +- 1.5 log,, CFU/g of gastric mucosa in normochlorhydric piglets sacrificed after 2 days to 6.9 +- 0.5 log-10 CFU/g in normochlorhydric piglets sacrificed after 5 days. Strain N6ureG::Km did not colonize any of seven normochlorhydric piglets and was recovered only in low numbers (lt 100 CFU/g) from four of nine achlorhydric piglets. the second experiment, piglets were coinoculated with both strains N6 and N6ureG::Km. Coinoculation did not affect colonization by urease-positive ***Urease*** - ***deficient*** N6ureG::Km was unable to colonize even in the presence of urease-positive ***bacteria*** These results confirm that urease enzymatic activity (and not urease protein) is essential for colonization, that this effect is independent of diffusible products of urea metabolism, and that gastric pH protection is not a major role of urease in promoting colonization by H. pylori.
- AB. . . the second experiment, piglets were coinoculated with both strains N6 and N6ureG::Km. Coinoculation did not affect colonization by urease-positive N6. ***Urease*** ***deficient*** N6ureG::Km was unable to colonize even in the presence of urease-positive ***bacteria*** . These results confirm that urease enzymatic activity (and not urease protein) is essential for colonization, that this effect is independent. . .

ORGN Classifier

Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms

```
Organism Name
        aerobic helical or vibrioid gram-negative ***bacteria***
        Helicobacter pylori
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
       Suidae
                85740
     Super Taxa
       Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia
       pig
     Taxa Notes
       Animals, Artiodactyls, . . .
L11 ANSWER 13 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
AN
     1993:333419 BIOSIS <<LOGINID::20080330>>
DN
    PREV199345028144
                  ***urease*** - ***deficient*** mutant of Helicobacter
ΤI
    An isogenic
    pylori colonizes gastric epithelial explants, but not germ-free piglets.
ΑU
     Eaton, K. A. [Reprint author]; Labigne, A. F.; Krakowka, S.
    Dep. Vet. Pathobiol., Ohio State Univ., Columbus, OH, USA
CS
    Gastroenterology, (1993) Vol. 104, No. 4 SUPPL., pp. A694.
SO
    Meeting Info.: 94th Annual Meeting of the American Gastroenterological
    Association. Boston, Massachusetts, USA. May 15-21, 1993.
    CODEN: GASTAB. ISSN: 0016-5085.
DT
    Conference; (Meeting)
LA
    English
    Entered STN: 16 Jul 1993
ED
     Last Updated on STN: 31 Aug 1993
                 ***urease*** - ***deficient***
                                                     mutant of Helicobacter
    An isogenic
     pylori colonizes gastric epithelial explants, but not germ-free piglets.
ORGN Classifier
       Aerobic Helical or Vibrioid Gram-Negatives
     Super Taxa
       Eubacteria; ***Bacteria***; Microorganisms
     Organism Name
        aerobic helical or vibrioid gram-negative ***bacteria***
        Helicobacter pylori
     Taxa Notes
            ***Bacteria*** , Eubacteria, Microorganisms
ORGN Classifier
        Suidae 85740
        Artiodactyla; Mammalia; Vertebrata; Chordata; Animalia
     Organism Name
        Suidae
     Taxa Notes
       Animals, Artiodactyls,. . .
L11 ANSWER 14 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on
     STN
ΑN
    1992:327557 BIOSIS <<LOGINID::20080330>>
DN
    PREV199294029398; BA94:29398
ΤI
    CHARACTERIZATION OF HELICOBACTER-PYLORI UREASE MUTANTS.
ΑIJ
    SEGAL E D [Reprint author]; SHON J; TOMPKINS L S
CS
     DEP MICROBIOL IMMUNOL, DIGESTIVE DISEASES CENTER, STANFORD UNIV, STANFORD,
```

CALIF 94305, USA

SO Infection and Immunity, (1992) Vol. 60, No. 5, pp. 1883-1889. CODEN: INFIBR. ISSN: 0019-9567.

DT Article

FS BA

LA ENGLISH

ED Entered STN: 11 Jul 1992 Last Updated on STN: 11 Jul 1992

The association between Helicobacter pylori, gastritis, and peptic ulcer AB is well established, and the association of infection with gastric cancer has been noted in several developing countries. However, the pathogenic mechanism(s) leading to disease states has not been elucidated. The H. pylori urease is thought to be a determinant of pathogenicity, since the enzyme is produced by all H. pylori clinical isolates. Evidence indicates that some H. pylori strains are more cytotoxic than others, with a correlation between the activity of the urease and the presence of a vacuolating cytotoxin having been made. However, the number of cytotoxins remains unknown at this time. The relationship between the urease and cytotoxicity has previously been examined with chemical inhibitors. To examine the role of the urease and its relationship to cytotoxicity, ***urease*** - ***deficient*** mutants were produced following ethyl methanesulfonate mutagenesis of H. pylori 87A300. Two mutants (the ure1 and ure5 mutants) which were entirely deficient in urease activity (Ure-) were selected. Characterization of the isolates at the protein level showed that the urease subunits lacked the ability to complex and form the active urease enzyme. The urel mutant was shown to be sensitive to the effects of low pH in vitro and exhibited no cytotoxicity to eucaryotic cells, whereas the parental strain (Ure+) produced a cytotoxic effect in the presence of urea. Interaction between the H. pylori Ure+ and Ureproduced a cytotoxic effect in the presence of urea. Interaction between the H. pylori Ure+ and Ure- strains and Caco-2 cells appeared to be ***bacterial*** types elicited pedestal formation similar in that both and actin condensation. These results indicate that the H. pylori ureas may have many functions, among them (i) protecting H. pylori against the acidic environment of the stomach, (ii) acting as a cytotoxin, with human gastric cells especially susceptible to its activity, and (iii) disrupting cell tight junctions in such a manner than the cells remain viable but an

AB. . . cytotoxicity has previously been examined with chemical inhibitors. To examine the role of the urease and its relationship to cytotoxicity, ***urease*** - ***deficient*** mutants were produced following ethyl methanesulfonate mutagenesis of H. pylori 87A300. Two mutants (the urel and ure5 mutants) which were. . . urea. Interaction between the H. pylori Ure+ and Ure- strains and Caco-2 cells appeared to be similar in that both ***bacterial*** types elicited pedestal formation and actin condensation. These results indicate that the H. pylori ureas may have many functions, among. . .

ORGN Classifier

Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa

Eubacteria; ***Bacteria*** ; Microorganisms
Taxa Notes

ionic flow between the cells occurs.

Bacteria , Eubacteria, Microorganisms

ORGN Classifier

Vertebrata 85150

Super Taxa

Chordata; Animalia

Taxa Notes

Animals, Chordates, Nonhuman Vertebrates, Vertebrates

- L11 ANSWER 15 OF 20 BIOSIS COPYRIGHT (c) 2008 The Thomson Corporation on ΑN 1992:41708 BIOSIS <<LOGINID::20080330>> PREV199242017858; BR42:17858 TT CONSTRUCTION OF ***UREASE*** ***DEFICIENT*** MUTANTS OF HELICOBACTER-PYLORI BY ALLELIC EXCHANGE. ΑU FERRERO R [Reprint author]; CUSSAC V; COURCOUX P; LABIGNE A CS UNITE ENTEROBACTERIES, INSERM U199, INST PASTEUR, 75724 PARIS CEDEX 15, FR SO Microbial Ecology in Health and Disease, (1991) Vol. 4, No. SPEC. ISSUE, pp. S136. Meeting Info.: VITH INTERNATIONAL WORKSHOP ON CAMPYLOBACTER HELICOBACTER AND RELATED ORGANISMS, SYDNEY, NEW SOUTH WALES, AUSTRALIA, OCTOBER 7-10, 1991. MICROB ECOL HEALTH DIS. ISSN: 0891-060X. DT Conference; (Meeting) FS BR LA ENGLISH Entered STN: 7 Jan 1992 ED Last Updated on STN: 8 Jan 1992 ***DEFICIENT*** MUTANTS OF CONSTRUCTION OF ***UREASE*** ΤI HELICOBACTER-PYLORI BY ALLELIC EXCHANGE. ORGN Classifier Aerobic Helical or Vibrioid Gram-Negatives 06210 Super Taxa Eubacteria; ***Bacteria*** ; Microorganisms Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms ORGN Classifier Enterobacteriaceae 06702 Super Taxa Facultatively Anaerobic Gram-Negative Rods; Eubacteria; ***Bacteria*** ; Microorganisms Taxa Notes ***Bacteria*** , Eubacteria, Microorganisms L11 ANSWER 16 OF 20 CABA COPYRIGHT 2008 CABI on STN 95:23391 CABA <<LOGINID::20080330>> AN 19941908449 DN Hydrogenase and urease in cyanobacterial photosynthesis and nitrogen TΤ fixation ΑU Ewart, G. D.; Mackerras, A. H.; Smith, G. D.; Kashyap, A. K. [EDITOR]; Kumar, H. D. [EDITOR] CS Department of Biochemistry, Fculty of Science, Australian National University, Canberra, ACT 2601, Australia. SO Recent advances in phycology, (1994) pp. 21-30. 26 ref. Publisher: Rastogi Publications. Meerut ISBN: 81-85711-05-4 CY India
- DT Miscellaneous
- LA English
- ED Entered STN: 1 Feb 1995 Last Updated on STN: 1 Feb 1995
- AB In the cyanobacterium Anabaena cylindrica both hydrogenase and urease activities are dependent on the presence of Ni in the growth medium. In

cyanobacteria there are two forms of hydrogenase: soluble and membrane bound. Electrophoretic analysis showed that the enzyme is a dimer consisting of 2 subunits. Tritium exchange and reductive hydrogenase activities could be differentially inhibited. Growth of cells in the absence of Ni produced hydrogenase and ***urease*** - ***deficient*** cells. The exponential growth rate of nitrogen-fixing cells in A. cylindrica was not inhibited by the absence of Ni. Growth of A. cylindrica was dependent on Ni when non-nitrogen-fixing cells were used to reinitiate nitrogen-fixing growth. Nickel-deficient cells showed a pronounced growth lag which was associated with loss of pigment, delayed nitrogenase synthesis, and cyanophycin accumulation. These observations suggested a role for Ni in nitrogen metabolism in addition to that as a cofactor for urease.

AB . . . exchange and reductive hydrogenase activities could be differentially inhibited. Growth of cells in the absence of Ni produced hydrogenase and ***urease*** - ***deficient*** cells. The exponential growth rate of nitrogen-fixing cells in A. cylindrica was not inhibited by the absence of Ni. Growth. . .

ORGN ***bacteria*** ; Cyanobacteria

- L11 ANSWER 17 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN
- AN 2006:380705 CAPLUS <<LOGINID::20080330>>
- DN 144:410795
- TI Recombinant ***Mycobacterium*** BCG adjuvant in vaccination
- IN Laeufer, Albrecht; Eisele, Bernd; Grode, Leander
- PA Vakzine Projekt Management G.m.b.H., Germany
- SO Eur. Pat. Appl., 17 pp.
 - CODEN: EPXXDW
- DT Patent
- LA English
- FAN CNT 1

FAN.CNT 1 PATENT NO.				KIND DATE			APPLICATION NO.					DATE							
ΡI	EP	 EP 1649869				A1 20060426			EP 2004-25096					20041021					
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙΤ,	LI,	LU,	NL,	SE,	MC,	PT,	
			IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	ВG,	CZ,	EE,	HU,	PL,	SK,	HR
	AU	2005	2989	76		A1		20060504		AU 2005-298976					20051016				
	CA	2584	2584321 2006045468			A1 20060504				CA 2005-2584321					20051016				
	WO	2006				A1 20060504				WO 2005-EP11127					20051016				
		W:	ΑE,	AG,	AL,	ΑM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,	
			CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	
			GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KM,	KΡ,	KR,	KΖ,	
			LC,	LK,	LR,	LS,	LT,	LU,	LV,	LY,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	
			NA,	NG,	NΙ,	NO,	NΖ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	
			SK,	SL,	SM,	SY,	ΤJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	
			YU,	ZA,	ZM,	ZW													
		RW:	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,	FΙ,	FR,	GB,	GR,	HU,	ΙE,	
			IS,	ΙΤ,	LT,	LU,	LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	
			CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	ΤG,	BW,	GH,	
			GM ,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑM,	ΑZ,	BY,	
			,	,	,	RU,	,												
	EP	1802340								EP 2005-795016					20051016				
		R:	,		,	,		,	,	,	EE,	,	,	,	,	,	,	IE,	
			•	,		,	•	,	,	,	PL,	•		,		,			
		IN 2007DN02871								CN 2005-80036326									
									IN 2007-DN2871										
	MX	200704734				А		2007	0713		MX 2007-4734					20070419			

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KR 2007068398 A 20070629 KR 2007-709076 20070420 PRAI EP 2004-25096 A 20041021
    WO 2005-EP11127 W
                              20051016
     The authors disclose the use of ***urease*** - ***deficient***
AΒ
      ***Mycobacterium*** BCG expressing listeriolysin as an adjuvant in
     vaccination. In one example, a tumor vaccine comprises a allogeneic
     prostate carcinoma cell line, transgenic for interferon-.gamma. and
     interleukin-2, in combination with the foregoing ***bacterial***
                                                                        cell
     adjuvant.
RE.CNT 6
             THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
     Recombinant ***Mycobacterium*** BCG adjuvant in vaccination
ΤI
     The authors disclose the use of ***urease*** - ***deficient***
AΒ
       ***Mycobacterium*** BCG expressing listeriolysin as an adjuvant in
     vaccination. In one example, a tumor vaccine comprises a allogeneic
     prostate carcinoma cell line, transgenic for interferon-.gamma. and
     interleukin-2, in combination with the foregoing ***bacterial*** cell
     adjuvant.
     ***Mycobacterium*** cytolysin adjuvant vaccine
ST
ΙT
        (antimalarial; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant for)
ΙT
    Antigens
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (autoantigens, microbial; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination against)
ΙT
     Prostate gland, neoplasm
        (carcinoma; ***urease*** - ***deficient*** ***Mycobacterium***
        BCG expressing listeriolysin as vaccine adjuvant for
       cytokine-transgenic cell immunogens)
ΙT
     Intestine, neoplasm
        (colon, carcinoma; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΙT
     Carcinoma
                ***urease*** - ***deficient***
                                                    ***Mycobacterium***
       BCG expressing listeriolysin as vaccine adjuvant for
       cytokine-transgenic cell immunogens)
ΙT
     Carcinoma
        (head and neck squamous cell carcinoma; ***urease*** -
                           ***Mycobacterium*** BCG expressing listeriolysin
          ***deficient***
        as vaccine adjuvant for cytokine-transgenic cell immunogens)
    Cell adhesion molecules
     Interleukin 12
     Interleukin 2
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (in combination with ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
ΙT
    Hemolysins
     RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (listeriolysins O; ***urease*** - ***deficient***
          ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
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ΙT
    Antigens
    Tumor antigens
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (microbial; ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant in vaccination against)
ΙT
    Lung, neoplasm
       (non-small-cell carcinoma;
                                 ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΙT
    Lysosome
       (phagolysosome; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination in relation to)
ΙT
    Carcinoma
       (prostatic; ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as vaccine adjuvant for
       cytokine-transgenic cell immunogens)
ΙT
    Carcinoma
       (pulmonary non-small-cell; ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΤT
    Kidney, neoplasm
                             ***urease*** - ***deficient***
       (renal cell carcinoma;
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΤТ
    Carcinoma
       (renal cell; ***urease*** - ***deficient*** ***Mycobacterium***
       BCG expressing listeriolysin as vaccine adjuvant for
       cytokine-transgenic cell immunogens)
ΙT
    Head and Neck, neoplasm
                                 ***urease*** - ***deficient***
       (squamous cell carcinoma;
         ***Mycobacterium*** BCG expressing listeriolysin as vaccine adjuvant
       for cytokine-transgenic cell immunogens)
ΙT
    Vaccines
       (tumor; ***urease*** - ***deficient***
                                                   ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant for)
ΙT
    MSP-1 (protein)
    RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
       BCG
       expressing listeriolysin as adjuvant for)
ΤT
    Plasmodium falciparum
       expressing listeriolysin as adjuvant for merozoite surface protein of)
ΙT
    Malaria
       ( ***urease*** - ***deficient***
                                           ***Mycobacterium*** BCG
       expressing listeriolysin as adjuvant for vaccination against)
ΙT
    Human
        ***Mycobacterium***
                           BCG
        ***Mycobacterium***
                            ***bovis***
       ( ***urease*** - ***deficient***
                                           ***Mycobacterium***
                                                                 BCG
       expressing listeriolysin as adjuvant in vaccination)
ΤT
    Antigen-presenting cell
    Brain, neoplasm
    Dendritic cell
    Mammary gland, neoplasm
    Melanoma
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Neoplasm
       expressing listeriolysin as vaccine adjuvant for cytokine-transgenic
       cell immunogens)
ΤТ
    Antimalarials
    Antitumor agents
        (vaccines; ***urease*** - ***deficient***
                                                       ***Mycobacterium***
       BCG expressing listeriolysin as adjuvant for)
ΙT
    Interferons
    RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (.gamma.; in combination with
                                     ***urease*** - ***deficient***
         ***Mycobacterium*** BCG expressing listeriolysin as adjuvant in
       vaccination)
ΤТ
    884349-82-0
    RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (amino acid sequence; ***urease*** - ***deficient***
         ***Mycobacterium***
                             BCG expressing listeriolysin as adjuvant in
       vaccination)
    9002-13-5D, Urease, subunit C
ΙT
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
                    ***urease*** - ***deficient***
                                                          ***Mvcobacterium***
        (deficiency;
       BCG expressing listeriolysin as adjuvant in vaccination)
    884349-81-9, DNA (Listeria monocytogenes gene hyl)
ΤТ
    RL: BSU (Biological study, unclassified); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (nucleotide sequence; ***urease*** - ***deficient***
                             BCG expressing listeriolysin as adjuvant in
         ***Mycobacterium***
       vaccination)
L11 ANSWER 18 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN
    1997:498189 CAPLUS <<LOGINID::20080330>>
DN
    127:188074
TΙ
    Interactions of a catalase- and an urease-negative mutant of Helicobacter
    pylori with polymorphonuclear granulocytes
    Marxer, Martin; Farzam, Fardad; Spiegelhalder, Christiane; Kersten,
ΑU
    Astrid; Odenbreit, Stefan; Haas, Rainer; Kist, Manfred
CS
    Inst. fur Med. Mikrobiologie und Hygiene, Freiburg, 79104, Germany
SO
    Campylobacters, Helicobacters, and Related Organisms, [Proceedings of the
    International Workshop on Campylobacters, Helicobacters, and Related
    Organisms], 8th, Winchester, UK, July 10-13, 1995 (1996), Meeting Date
    1995, 701-705. Editor(s): Newell, Diane G.; Ketley, Julian M.; Feldman,
    Roger A. Publisher: Plenum, New York, N. Y.
    CODEN: 64TNAY
    Conference
DT
LA
    English
    To examine whether or not catalase and urease play a role as virulence
AB
    factors of H. pylori, isogenic catalase- or ***urease*** -
      ***deficient*** mutant strains, constructed by transposon mutagenesis,
    were compared with the corresponding wild-type strain 69A with respect to
    their interactions with polymorphonuclear nucleophiles (PMNs), including
    sensitivity towards killing by PMNs, strength of the oxidative burst, and
    electron microscopic studies. The results from the the catalase-neq.
    mutant indicated that although catalase is able to scavenge hydrogen
    peroxide, it does not protect the ***bacteria*** efficiently from
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PMN-induced killing. In the case of the urease-neg. mutant, the phagocytic oxidative burst in the presence of the mutant was not significantly increased compared to that induced by the wild type, thus suggesting that non-oxygen-mediated killing mechanisms of the PMNs are responsible form the more efficient ***bactericidal*** activity on the ***urease*** - ***deficient*** mutant. To examine whether or not catalase and urease play a role as virulence factors of H. pylori, isogenic catalase- or ***urease*** -***deficient*** mutant strains, constructed by transposon mutagenesis, were compared with the corresponding wild-type strain 69A with respect to their interactions with. . . from the the catalase-neg. mutant indicated that although catalase is able to scavenge hydrogen peroxide, it does not protect the ***bacteria*** efficiently from PMN-induced killing. In the case of the urease-neg. mutant, the phagocytic oxidative burst in the presence of the. . . induced by the wild type, thus suggesting that non-oxygen-mediated killing mechanisms of the PMNs are responsible form the more efficient ***bactericidal*** activity on the ***urease*** - ***deficient*** mutant. L11 ANSWER 19 OF 20 CAPLUS COPYRIGHT 2008 ACS on STN 1992:172374 CAPLUS <<LOGINID::20080330>> 116:172374 Selection of L-lysine-producing strain Au111-2 Su, Lingming; Xu, Suowei; Fu, Yinghua; Wang, Xingzhen; Tang, Shanghua; Shao, Guoliang Shanghai Inst. Ind. Microbiol., Shanghai, Peop. Rep. China Gongve Weishengwu (1991), 21(6), 12-16 CODEN: GOWEEK; ISSN: 1001-6678 Journal Chinese ***Bacteria*** strain All1 was a good producer of lysine, but was ***urease*** ***deficient*** , and so the pH in the process of fermn. could not be controlled with urea. After the mutation with MNNG and screening with urea as nitrogen source, an urease revertant strain Aul11-2 was obtained. The lysine productivity and the conversion ratio to the glucose of the urease revertant Au111-2 increased by 25% and 15% than that of strain All1. ***Bacteria*** strain A111 was a good producer of lysine, but was ***deficient*** , and so the pH in the process of ***urease*** fermn. could not be controlled with urea. After the mutation with MNNG. . . lysine fermn ***bacteria*** urease ***Bacteria*** (lysine formation by, urease mutation effect on) Fermentation ***bacteria*** , urease mutation effect on) (lysine, with 56-87-1, L-Lysine, biological studies RL: FORM (Formation, nonpreparative) (formation of, by ***bacteria*** , urease mutation effect on) 9002-13-5, Urease RL: BIOL (Biological study) (of ***bacteria*** , lysine formation in relation to) L11 ANSWER 20 OF 20 MEDLINE on STN 2007476473 MEDLINE <<LOGINID::20080330>>

ΑN DN

ΤI

ΑU

CS

SO

DТ

LA

AΒ

AB

ST

ΙT

ΙT

ΤТ

ΙT

AN

DN

PubMed ID: 17519853

- TI [Strategies for the development of new ***tuberculosis*** vaccines].

 Strategie per lo sviluppo di nuovi vaccini antitubercolari.
- AU Fattorini L
- CS Dipartimento di Malattie Infettive, Parassitarie e Immunomediate, Istituto Superiore di Sanita, Roma, Italy.. lanfranco.fattorini@iss.it
- SO Minerva medica, (2007 Apr) Vol. 98, No. 2, pp. 109-19. Ref: 47 Journal code: 0400732. ISSN: 0026-4806.
- CY Italy
- DT (ENGLISH ABSTRACT)
 Journal; Article; (JOURNAL ARTICLE)
 General Review; (REVIEW)
- LA Italian
- FS Priority Journals
- EM 200708
- ED Entered STN: 16 Aug 2007 Last Updated on STN: 17 Aug 2007 Entered Medline: 16 Aug 2007
- ***Tuberculosis*** remains a substantial global health problem causing 2 million deaths, and an estimated 8 to 10 million new infections a year. The efficacy of the ***Mycobacterium*** ***bovis*** Bacillus Calmette-Guerin (BCG), the only available antituberculosis vaccine, is variable (0-80%), especially in ***tuberculosis*** -endemic countries. Over the past decade there has been a resurgence of interest in the development of new ***tuberculosis*** vaccines and some of the most promising are now entering into early clinical trials, based on two different strategies. The first is to use whole ***mycobacteria*** to replace BCG (priming vaccines), either by developing a recombinant strain of BCG or an attenuated strain of ***Mycobacterium***
 - ***tuberculosis*** . To date, two recombinant strains of BCG, one overexpressing antigen 85B (rBCG-85B) and the other, a ***urease*** ***deficient*** BCG mutant which expresses the listeriolysin O gene

from

Listeria monocytogenes (rBCG::DeltaureC-hly+), entered into clinical trials. The second approach is to develop subunit vaccines (recombinant proteins and viral vectors, and DNA vaccines) expressing immunodominant antigen/s from M. ***tuberculosis*** able to augmenting BCG protection (booster vaccines). At the moment, three major vaccines, namely a recombinant modified vaccinia virus Ankara expressing antigen 85A (MVA85A), a fusion protein of ESAT6 and 85B (Hybrid 1), and another fusion protein comprising the 32 and 39 Kda proteins (72f) entered into clinical trials.

- TI [Strategies for the development of new ***tuberculosis*** vaccines].

 Strategie per lo sviluppo di nuovi vaccini antitubercolari.
- ***Tuberculosis*** remains a substantial global health problem causing 2 million deaths, and an estimated 8 to 10 million new infections a year. The efficacy of the ***Mycobacterium*** ***bovis*** Bacillus Calmette-Guerin (BCG), the only available antituberculosis vaccine, is variable (0-80%), especially in ***tuberculosis*** -endemic countries. Over the past decade there has been a resurgence of interest in the development of new ***tuberculosis*** vaccines and some of the most promising are now entering into early clinical trials, based on two different strategies. The first is to use whole ***mycobacteria*** to replace BCG (priming vaccines), either by developing a recombinant strain of BCG or an attenuated strain of ***Mycobacterium***

tuberculosis . To date, two recombinant strains of BCG, one overexpressing antigen 85B (rBCG-85B) and the other, a ***urease*** -

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***deficient*** BCG mutant which expresses the listeriolysin O gene
from
    Listeria monocytogenes (rBCG::DeltaureC-hly+), entered into clinical
    trials. The second approach is to develop subunit vaccines (recombinant
    proteins and viral vectors, and DNA vaccines) expressing immunodominant
     antiqen/s from M. ***tuberculosis*** able to augmenting BCG protection
     (booster vaccines). At the moment, three major vaccines, namely a
     recombinant modified vaccinia virus Ankara.
     Immunization, Secondary: MT, methods
CT
        ****Mycobacterium bovis: IM, immunology***
         ****Mycobacterium tuberculosis: IM, immunology***
         ****Tuberculosis Vaccines: IM, immunology***
     Vaccines, Synthetic: IM, immunology
CN
    0 ( ***Tuberculosis*** Vaccines); 0 (Vaccines, Synthetic)
```